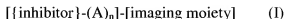


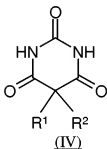
**Listing of Claims:**

1. (Currently amended) An *in vivo* imaging agent of Formula I:



where:

{inhibitor} is a synthetic barbituric acid matrix metalloproteinase inhibitor, of Formula IV, which is labeled at the R<sup>2</sup> substituent 5-position of the barbituric acid with said imaging moiety;



wherein:

R<sup>1</sup> is R" or a Z group;

R<sup>2</sup> is R", Y or -NR<sup>4</sup>R<sup>5</sup>, where R<sup>4</sup> is H or an R" group, R<sup>5</sup> is H, C<sub>2-14</sub> acyl, C<sub>2-10</sub> aminoalkyl or (N-C<sub>2-14</sub> acyl)C<sub>2-10</sub> aminoalkyl or an R" group, or R<sup>4</sup> and R<sup>5</sup> together with the N atom to which they are attached form an optionally (N-C<sub>2-14</sub>)acylated C<sub>2-8</sub> cycloaminoalkylene ring;

R" is independently C<sub>1-14</sub> alkyl, C<sub>3-8</sub> cycloalkyl, C<sub>2-14</sub> alkenyl, C<sub>1-14</sub> fluoroalkyl, C<sub>1-14</sub> perfluoroalkyl, C<sub>6-14</sub> aryl, C<sub>2-14</sub> heteroaryl or C<sub>7-16</sub> alkylaryl;  
Z is a group of formula -A<sup>1</sup>O(A<sup>2</sup>O)<sub>p</sub>R<sup>3</sup> where p is 0 or 1, and A<sup>1</sup> and A<sup>2</sup> are independently C<sub>1-10</sub> alkylene, C<sub>3-8</sub> cycloalkylene, C<sub>1-10</sub> perfluoroalkylene, C<sub>6-10</sub> arylene or C<sub>2-10</sub> heteroarylene, and R<sup>3</sup> is an R group where R is independently chosen from H, C<sub>1-4</sub> alkyl, C<sub>2-4</sub> alkenyl, C<sub>2-4</sub> alkynyl, C<sub>1-4</sub> alkoxyalkyl or C<sub>1-4</sub> hydroxyalkyl;

Y is a group of formula:



where E is CR<sub>2</sub>, O, S or NR<sup>6</sup>; and R<sup>6</sup> is C<sub>2-14</sub> acyl, or an R" or Z group;

-(A)<sub>n</sub>- is a linker group wherein each A is independently -CR<sub>2</sub>-, -CR=CR-,  
-C≡C-, -CR<sub>2</sub>CO<sub>2</sub>-, -CO<sub>2</sub>CR<sub>2</sub>-, -NRCO-, -CONR-, -NR(C=O)NR-,  
-NR(C=S)NR-, -SO<sub>2</sub>NR-, -NRSO<sub>2</sub>-, -CR<sub>2</sub>OCR<sub>2</sub>-, -CR<sub>2</sub>SCR<sub>2</sub>-,  
-CR<sub>2</sub>NRCR<sub>2</sub>-, a C<sub>4-8</sub> cycloheteroalkylene group, a C<sub>4-8</sub> cycloalkylene group, a  
C<sub>5-12</sub> arylene group, or a C<sub>3-12</sub> heteroarylene group, an amino acid or a  
monodisperse polyethyleneglycol (PEG) building block;

R is independently chosen from H, C<sub>1-4</sub> alkyl, C<sub>2-4</sub> alkenyl, C<sub>2-4</sub> alkynyl,  
C<sub>1-4</sub> alkoxyalkyl or C<sub>1-4</sub> hydroxyalkyl;

n is an integer of value 0 to 10;

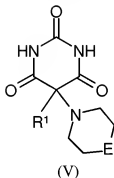
wherein the imaging moiety can be detected externally in a non-invasive  
manner following administration of said labelled synthetic barbituric acid  
matrix metalloproteinase inhibitor to the mammalian body *in vivo*, and said  
imaging moiety is chosen from:

- (i) a radioactive metal ion, which is a gamma emitter or a positron emitter  
and is chosen from <sup>99m</sup>Tc, <sup>111</sup>In, <sup>64</sup>Cu, <sup>67</sup>Cu, <sup>67</sup>Ga or <sup>68</sup>Ga;
- (ii) the gamma-emitting radioactive halogen <sup>123</sup>I;
- (iii) a positron-emitting radioactive non-metal chosen from <sup>18</sup>F, <sup>11</sup>C or <sup>13</sup>N.

2. (Cancelled)

3. (Previously presented) The imaging agent of Claim 1, where the synthetic barbituric acid  
matrix metalloproteinase inhibitor is conjugated to a ligand, and said ligand forms a metal  
complex with the radioactive metal ion.

4. (Original) The imaging agent of Claim 3, where the ligand is a chelating agent.
5. (Cancelled)
6. (Cancelled)
7. (Cancelled)
8. (Cancelled)
10. (Currently amended) The imaging agent of claim 19, where  $R^2$  is Y or  $-NR^4R^5$ .
11. (Cancelled)
12. (Currently amended) The imaging agent of claim 19, of Formula V:



where E is CHR or NR<sup>6</sup> and R<sup>1</sup> is C<sub>6-14</sub> *n*-alkyl, or C<sub>6-14</sub> aryl.

13. (Currently amended) The imaging agent of claim 12, where E is NR<sup>6</sup> and R<sup>6</sup> is C<sub>2-14</sub> acyl;  $-(CH_2)_dOH$ , where d is 2, 3, 4 or 5; or  $-C_6H_4X$ , where X is H, C<sub>1-4</sub> alkyl, Hal, OR, NR<sub>2</sub>, NO<sub>2</sub> or SO<sub>2</sub>NR<sup>7</sup>R<sup>8</sup>, where R<sup>7</sup> and R<sup>8</sup> are independently R groups, and R is as defined in Claim 19.
14. (Previously presented) The imaging agent of claim 12, where R<sup>1</sup> is *n*-octyl, *n*-decyl, biphenyl, C<sub>6</sub>H<sub>5</sub>X or  $-C_6H_4-O-C_6H_4X$  where X is as defined in Claim 13.

15. (Cancelled)

16. (Cancelled)

17. (Currently Amended) The radiopharmaceutical composition of claim 146, where the imaging moiety comprises a radioactive metal ion.

18. (Original) The radiopharmaceutical composition of claim 16, where the imaging moiety comprises a positron-emitting radioactive non-metal or a gamma-emitting radioactive halogen.

19. (Withdrawn) A conjugate of a synthetic barbituric acid matrix metalloproteinase inhibitor with a ligand, wherein the barbituric acid comprises a 5-position substituent, and said 5-position substituent comprises a ligand capable of forming a metal complex with a radioactive metal ion which is resistant to transchelation.

20. (Withdrawn) The conjugate of Claim 19, of Formula Ib:



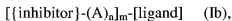
where {inhibitor}, A, n and m are as defined in Claim 2.

21. (Withdrawn) The conjugate of Claim 19, wherein the synthetic barbituric acid matrix metalloproteinase inhibitor is of Formula IV or Formula V of Claims 9 to 14.

22. (Withdrawn) The conjugate of Claim 19, wherein the ligand is a chelating agent.

23. (Withdrawn) The conjugate of Claim 22, wherein the chelating agent has a diaminedioxime,  $N_2S_2$ , or  $N_3S$  donor set.

24. (Withdrawn) A kit for the preparation of the radiopharmaceutical composition of Claim 17, which comprises a conjugate of a synthetic barbituric acid matrix metalloproteinase inhibitor with a ligand, wherein the barbituric acid comprises a 5-position substituent, and said 5-position substituent comprises a ligand capable of forming a metal complex with a radioactive metal ion which is resistant to transchelation, said conjugate being of Formula Ib:



where {inhibitor}, A, n and m are as defined in Claim 2, and wherein the ligand is a chelating agent.

25. (Withdrawn) The kit of Claim 26, where the radioactive metal ion is  $^{99m}\text{Tc}$ , and the kit further comprises a biocompatible reductant.

26. (Withdrawn) A kit for the preparation of the radiopharmaceutical composition of Claim 18, which comprises a precursor in sterile form which is a non-radioactive derivative of the barbituric acid matrix metalloproteinase inhibitor of claims 1, wherein said non-radioactive derivative is capable of reaction with a source of the positron-emitting radioactive non-metal or gamma-emitting radioactive halogen to give the desired radiopharmaceutical.

27. (Original) The kit of Claim 26, where the source of the positron-emitting radioactive non-metal or gamma-emitting radioactive halogen is chosen from:

- (i) halide ion;
- (ii)  $\text{F}^+$  or  $\text{I}^+$ ; or
- (iii) an alkylating agent chosen from an alkyl or fluoroalkyl halide, tosylate, triflate or mesylate;
- (iv)  $\text{HS}(\text{CH}_2)_3^{18}\text{F}$ .

28. (Previously presented) The kit of claim 26, wherein the non-radioactive derivative is chosen from:

- (i) an organometallic derivative such as a trialkylstannane or a trialkylsilane;

- (ii) a derivative containing an alkyl or aryl iodide or bromide, alkyl tosylate or alkyl mesylate for nucleophilic substitution;
  - (iii) a derivative containing an aromatic ring activated towards nucleophilic or electrophilic substitution;
  - (iv) a derivative containing a functional group which undergoes facile alkylation;
  - (v) a derivative which undergoes alkylation with an alkyl thiol to give a thioether.
29. (Previously presented) The kit of claim 26, where the precursor is bound to a solid phase.
30. (Withdrawn) Use of the imaging agent of Claim 1 for the diagnostic imaging of atherosclerosis.
31. (Withdrawn) Use of the imaging agent of Claim 1 for the diagnostic imaging of unstable plaques.
32. (Withdrawn) Use of the imaging agent of Claim 1 for the intravascular detection of atherosclerosis.